

GenCore version 4.5
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OM protein - protein search, using sw model

Run on: November 5, 2001, 15:20:30 ; Search time 65.26 Seconds
(without alignments)
341.858 Million cell updates/sec

Title: US-09-593-316-4

Perfect score: 2008
Sequence: I MANVKKIVLISMIVSTIVV.....IKLVKMSQTKENYVKNV 368

Scoring table:
BLOSUM62
Gapop 10.0, Gapext 0.5

Searched: 412676 seqs, 60623988 residues

Total number of hits satisfying chosen parameters: 412676

Minimum DB seq length: 0
Maximum DB seq length: 2000000000

Post-processing: Minimum Match 08
Maximum Match 1008

Listing first 45 summaries

Database :

```

1: A.Geneseq.0601:*
2: /cgnl_9/gcqdada/geneseq/geneseq/AA1980.DAT:*
3: /cgnl_9/gcqdada/geneseq/geneseq/AA1981.DAT:*
4: /cgnl_9/gcqdada/geneseq/geneseq/AA1982.DAT:*
5: /cgnl_9/gcqdada/geneseq/geneseq/AA1983.DAT:*
6: /cgnl_9/gcqdada/geneseq/geneseq/AA1984.DAT:*
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16: /cgnl_9/gcqdada/geneseq/geneseq/AA1995.DAT:*
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19: /cgnl_9/gcqdada/geneseq/geneseq/AA1999.DAT:*
20: /cgnl_9/gcqdada/geneseq/geneseq/AA2000.DAT:*
21: /cgnl_9/gcqdada/geneseq/geneseq/AA2001.DAT:*
22: /cgnl_9/gcqdada/geneseq/geneseq/AA2001.DAT:*

```

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	1711.5	85.2	375	19	AAW49686
2	1707.5	85.0	371	16	AAW85082
3	1707	85.0	376	16	AAW80016
4	1700.5	84.7	371	16	AAW67777
5	1648.5	82.1	359	15	AAW62508
6	1648.5	82.1	359	17	AAW90573
7	1646.5	82.0	363	19	AAW49687
8	1604	79.9	354	19	AAW49688
9	1543	76.8	342	19	AAW49689
10	1476.5	73.5	394	12	AAW13750
11	1476.5	73.5	394	15	AAW45935

12	1476.5	73.5	394	18	AAW13639	Murine alpha(1,3)-galactosyl transferase
13	1326	66.0	313	15	AAW62507	Human A transferase
14	658	32.8	375	15	AAW57021	Human A transferase
15	656	32.7	353	12	AAW11317	Human A transferase
16	656	32.7	353	12	AAW57011	Human A transferase
17	655	32.6	354	12	AAW11789	Human A transferase
18	655	32.6	354	12	AAW11790	Human A transferase
19	655	32.6	354	12	AAW11792	Human A transferase
20	655	32.6	354	12	AAW57020	Human A transferase
21	641	31.9	358	15	AAW57013	Human A transferase
22	639	31.8	354	15	AAW57013	Human A transferase
23	639	31.8	354	15	AAW57016	Human A transferase
24	473	23.6	195	15	AAW57024	Human A transferase
25	271.5	13.5	100	20	AAW26039	Secreted protein
26	266.5	13.3	106	21	AAW86491	Human gene 59-enc
27	253	12.6	154	15	AAW57025	Human gene 59-enc
28	224.5	11.2	90	21	AAW86492	Human secreted pro
29	224.5	11.2	90	21	AAW86493	Human secreted pro
30	224.5	11.2	90	21	AAW86489	Human gene 59-enc
31	102.5	5.1	517	16	AAW66402	GalNAc-transferase
32	102.5	5.1	517	16	AAW16489	Honeybee mellitin
33	102.5	5.1	559	16	AAW66397	Cattle GalNAc-tran
34	102.5	5.1	559	16	AAW66401	GalNAc-transferase
35	102.5	5.1	559	18	AAW16484	Bovine N-acetylgl
36	101.5	5.1	561	21	AAW43561	Human cancer assoc
37	95	4.7	703	22	AAW70687	Pyrococcus furiosu
38	95	4.7	722	22	AAW70686	Pyrococcus furiosu
39	92.5	4.6	246	16	AAW74036	Human chondrocalci
40	92.5	4.6	1418	21	AAW96124	Collagen type II a
41	92.5	4.6	1487	19	AAW61562	Human type II coll
42	90	4.5	708	20	AAW06547	SV40 large T anti
43	89.5	4.5	1418	15	AAW59751	Type II collagen
44	89.5	4.5	1418	16	AAW71703	Collagen alpha 1 (
45	89.5	4.5	1418	22	AAW35624	Human type II coll

ALIGNMENTS

RESULT 1	
AAW49686	AAW49686 standard; Protein; 375 AA.
XX	XX
AC	AAW49686:
XX	XX
DT	10-NOV-1998 (first entry)
XX	XX
DE	Porcine alpha-1,3-galactosyl transferase isoform 1.
XX	XX
KW	Isoform: porcine; enzyme: alpha-1,3-galactosyl transferase; galactose;
KW	sugar: N-acetyllactosamine; glycoprotein; glycolipid; antibody; pig;
XX	graft tissue rejection; organ transplantation; xenotransplant.
OS	Sus scrofa.
XX	XX
PN	FR2751346-A1.
XX	XX
PD	23-JAN-1998.
XX	XX
PE	19-JUL-1996: 96FR-0009077.
XX	XX
PR	19-JUL-1996: 96FR-0009077.
XX	XX
PA	(INRM) INSERM INST NAT SANTE & RECH MEDICALE.
XX	XX
PI	Pourcel C, Souillou JP, Vannoy B;
XX	XX
DR	WPI, 1998-112876/11.
XX	XX
PT	N-PDB: AAW49453.
XX	XX
PT	Transgenic non-human donors of organs for human recipients -
XX	containing DNA encoding antibodies that inhibit graft rejection

claim 4; Page 42-44; 71pp; French.

This sequence represents isoform 1 of the porcine enzyme alpha-1,3-galactosyl transferase (alpha-1,3-GT). The enzyme catalyses the attachment of a galactose sugar molecule on the N-acetyllactosamine moiety found on surface glycoproteins and glycolipids. These sugar molecules are partly responsible for raising anti-graft antibodies, which lead to graft tissue rejection. The invention relates to a method of inhibiting the graft rejection mechanism by introducing the sequence encoding an antibody targeted to alpha-1,3-GT into the cells of animal, especially a pig, from whom organs may be used for xenotransplants. Neutralisation of the alpha-1,3-GT leads to tissues or organs lacking the galactose on the glycoproteins and glycolipids, thus preventing induction of the rejection response.

Sequence 375 AA:

Query Match: 85.2%; Score 1711.5; DB 19; Length 375;
Best Local Similarity: 84.9%; Pred. No. 7,60-165;
Matches 312; Conservative 40; Mismatches 29; Indels 5; Gaps 4;

1 MNVKKVTLISMLVSTVIVVWEYIHSPGSLFWINPSKRPVGGSSLOKQKMLPKMFNN 60
|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
5 mvvkvivlsmllsvtlvvnvfwyinspslftwlygskpvcv-qssqgrqwfpswln 63
61 G---YHEEGDGLNEKKKRNED-EKKIKLSQWENPKRPVNTMTKAPVWEGTYNKA 116
|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
64 qhsyveedaiaquekqkshdngelplvadtprckprevtlckkarpvweqlynda 123
117 VUNYYAKOKITVGLTVPAVGNVLEHYLEPFTSANKHFMVGNVFTYIMVDVSRMPL 176
|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
124 vldnyyakklitvrltvavgytychyleelissanylmvghkvlfymvddisrmp 183
177 ELGPKSEKVERIKRPKKMODISMKMKKTGCHLVANHONEDPFCMIVGVDEQRKGV 246
|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
184 clqlplstkvtklskcktwqdismmrmtklqchllahqhwvdtlrmvdyvqgnlqv 243
247 ETLGGSVAQLQAWYKADINDFTYERKRSAAVTPGEGDFYUAAIFGGTGYVNLTO 296
|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
244 clqlpsvaqlqawykadndfttyerkesaaylprqgddlyhhaallqqlrqlvnlq 303
297 EFGKGLKQKKNDFEAMQNDESHINKVPLLNKDTLLSPYGCMDYHIGLPADIKLVKMSM 356
|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
304 cctkqllgkknldfcawideshlnkyltlmkprklispycwdyhlgmsvdlrtvklaw 363
457 QTKRYNVNRRNV 368
|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
464 qkkyvnlvrnml 375

RESULT 2

AAAB5082 standard; Protein: 371 AA.

AAAB5082:

04 JUN 1996 (first entry)

Porcine alpha (1,3) galactosyltransferase.
Transgenic swine: porcine alpha (1,3) galactosyltransferase;
antisense; ribozyme; Gal-alpha-1,3-Gal-beta-1-4GlcNAc; epilope;
terminal; xenogenic; transplant; rejection; gene therapy; pig.
SUS scrofa.

W09528412 A1.

26 OCT 1995

01 NOV 1995 9500-0850-0940.

XX

13-APR-1994: 9AUS-0228933.

(BIOT-) BIOTRANSPLANT INC.

(GENE-) GEN HOSPITAL CORP.

(CHILD-) INST CHILD HEALTH.

Haetscher MW, Gustafsson KT, Sachs DH;

WPI: 1995-373759/48.

N-PDB: AAT02892.

Novel transgenic alpha (1,3) galactosyltransferase negative swine

used to produce rejection resistant cells for xenogenic

transplantation

Claim 11: Pages 35-37; 56pp; English.

transgenic swine in which the normal expression of the alpha (1,3) galactosyltransferase (AGT) AAB5082 is prevented, are prepd. by inhibiting the expression of the AGT gene AAT02892 using antisense oligonucleotides or ribozyme inactivators in a pluripotent porcine embryonic stem cell. It is then inserted into a porcine oocyte (from which the pronuclear material has been removed), which is itself grown to produce the transgenic swine. Swine which do not express AGT will not produce carbohydrate molecules confg. the distinctive terminal Gal-alpha-1,3-Gal-beta-1-4GlcNAc epitope, which is a significant factor in xenogenic (esp. human) transplant rejection of swine grafts. Therefore the swine cells produced in the AGT negative transgenic swine are xenogenic transplant rejection resistant, and can therefore be used by a transplant recipient, or to provide gene therapy.

Sequence 371 AA:

Query Match: 85.0%; Score 1707.5; DB 16; Length 371;
Best Local Similarity: 83.9%; Pred. No. 1,90-164;
Matches 312; Conservative 29; Mismatches 26; Indels 5; Gaps 3;

1 MNVKKVTLISMLVSTVIVVWEYIHSPGSLFWINPSKRPVGGSSLOKQKMLPKMFNN 60
|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
1 mvvkvivlsmllsvtlvvnvfwyinspslftwlygskpvcv-qssqgrqwfpswln 59
61 G---YHEEGDGLNEKKKRNED-EKKIKLSQWENPKRPVNTMTKAPVWEGTYNKA 116
|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
60 qhsyveedaiaquekqkshdngelplvadtprckprevtlckkarpvweqlynda 119
117 VUNYYAKOKITVGLTVPAVGNVLEHYLEPFTSANKHFMVGNVFTYIMVDVSRMPL 176
|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
120 vldnyyakklitvrltvavgytychyleelissanylmvghkvlfymvddisrmp 179
177 ELGPKSEKVERIKRPKKMODISMKMKKTGCHLVANHONEDPFCMIVGVDEQRKGV 246
|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
180 clqlplstkvtklskcktwqdismmrmtklqchllahqhwvdtlrmvdyvqgnlqv 239
247 ETLGGSVAQLQAWYKADINDFTYERKRSAAVTPGEGDFYUAAIFGGTGYVNLTO 296
|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
240 clqlpsvaqlqawykadndfttyerkesaaylprqgddlyhhaallqqlrqlvnlq 299
297 EFGKGLKQKKNDFEAMQNDESHINKVPLLNKDTLLSPYGCMDYHIGLPADIKLVKMSM 356
|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
300 cctkqllgkknldfcawideshlnkyltlmkprklispycwdyhlgmsvdlrtvklaw 359
357 QTKRYNVNRRNV 368
|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
360 qkkyvnlvrnml 371

RESULT 3

AAAB0016 standard; Protein: 376 AA.

AAAB0016:

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XX 01-MAY-1996 (first entry)
DT
XX Marmoset alpha-1,3-galactosyltransferase.
XX
DE Marmoset; alpha-1,3-galactosyltransferase; immune response; glycoprotein;
KM alpha-galactosyl epitope; cell membrane; virus; phagocytosis; tumour;
KW antigen processing; leukemia; lymphoma; myeloma; melanoma; carcinoma;
XX sarcoma; vaccine; opsonisation; glycoprotein; antibody; anti-Gal.
XX
OS Callithrix jacchus.
XX
XX W09524924-A1.
XX
FN 21-SEP-1995.
PD
XX 13-MAR-1995; 95WO-0503156.
XX
PR 15-MAR-1994; 94US-0213200.
XX
XX (UYHA-) UNIV HAHNEMANN & MEDICAL COLLEGE PENNSYL.
XX
XX Galili U, Repik PM;
XX
XX WPI: 1995-336816/43.
XX
DR N-PSDB: MAT04522.
XX
PT Association of an alpha-galactosyl epitope with a tumour or viral
PT antigen - is administered to anti-Gal synthesising animals to induce
PT an immune response
XX
XX Disclosure: Fig 9; 85pp; English.
XX
XX The amino acid sequence of the marmoset alpha-1,3-galactosyltransferase,
XX CC The enzyme can be used in methods of enhancing an immune response by
XX CC associating the alpha-galactosyl epitope with a cell membrane or viral
XX CC glycoprotein. The alpha-galactosyl epitope enhances phagocytosis and
XX CC subsequent processing of the antigen. The method is useful in the
XX CC treatment of tumours e.g. leukemia, lymphoma, myeloma, melanoma,
XX CC carcinoma and sarcoma, or for the generation of viral vaccines by
XX CC opsonising a viral glycoprotein. The alpha-galactosyl epitope enhances
XX CC recognition of the antigen in an animal that synthesises the naturally
XX CC occurring antibody - anti-Gal.
XX
XX Sequence 376 AA:
SQ
Query Match 85.0%; Score 1707; DB 16; Length 376;
Best Local Similarity 82.2%; Pred. No. 2.2e-164;
Matches 309; Conservative 33; Mismatches 26; Indels 8; Gaps 2;
QY 1 MNVKGKVLISMLVSTVIVFMEVYTHSEGSILFWINPSKNEPVGSSIOKGMLEPRMENN 60
DB 1 mnykvkvlismlvstvlvltwelyinspegsilwlyhsknpevdssaqkdwfwgwlfn 60
QY 61 GYH-----EEDGDI---NEEKQÖRNEDESKKLKLSMFPNFKREEVYMTKMAPVWVEST 112
DB 61 gihnygqecddtckekyreeegkckeddtelrlwdwfnppkkrpevmvtvqwkapyvwegt 120
QY 113 YNRAYLWDWYVKKOKITVCLTVFAYGORYLEHYLEEFLLTANKIKFMVGHPIYFIIMDDVSR 172
DB 121 ynkalenyvayqkltvcltvclalqirylehyleeftvlsanryfwmqhkvtlymwddvsk 180
QY 173 MPELEIEMLSKFKYERIKPEKRMQDISMKRMKTTIDEHIVAHQHEHVDLFEGCDVQVQFOD 232
DB 181 apilelgelstikvlevrepekrwqslsmmrkljgenllahlghevdltfcmadvqavtgd 240
QY 233 KFGVETLGEASVQIQAMWKADPNDFTFERRKESAAVIRFEGSDYVYUAAIFGQIFPTOVL 292
DB 241 hfyvetlqgsvaqlqawwykadpndftfyerkrasaaylirfgqdflynaatfgstprlgl 300
QY 293 NTGCGFGLILDKKKNQDIEAQWHDESHLNKVFLLKKPTKILSPKVCWYHGLDPLADIKLY 352

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Db 301 ntgcctcgllldkndleewesheshnkyl1lnkpskllspsycwadyhquqpsdtklv 360
QY 353 KMSWOTKEYNNVNRNV 368
   1:|||||:||||
Db 361 kswgtkeynlvrkxv 376

RESULT 4
ID AAR76777 standard; Protein: 371 AA.
AC AAR76777:
DT 11-DEC-1995 (first entry)
DE Pig alpha-1,3-galactosyltransferase.
KW Alpha-1,3-galactosyltransferase; alpha-1,3-galT; transgenic animal;
KW pig; hyperacute rejection; xenotransplantation; donor organ;
KW allograft rejection; Gal epitope; gene disruption;
KW homologous recombination; knock-out.
OS Sus scrofa.
XX
XX W09520661-A1.
XX
XX 03-AUG-1995.
XX
XX 27-JAN-1995; 95W0-1B00088.
XX
XX 26-JAN-1995; 95US-0188607.
XX 27-JAN-1994; 94US-0188607.
XX
XX (BRES-) BRESATPC LTD.
XX (SVIN-) ST VINCENT'S HOSPITAL, MELBOURNE LTD.
XX
XX Crawford RJ, Dapice AJF, Pearse MJ, Rathjen PD;
XX Robbins AJ;
XX
XX WPI: 1995-275446/36.
XX N-PSDR; AAO93077.
XX
XX New alpha-1,3-galactosyltransferase and leukaemia inhibitor factor
XX - corresp. DNA and nucleic acid constructs for inactivating the
XX transplants gene; for eliminating hyperacute region in human
XX transplants
XX
XX Claim 3; Fig.5: 184pp; English.
XX
XX cDNA encoding porcine alpha-1,3-galT was generated from liver RNA
XX using primers based on conserved regions of the mouse and cattle alpha-
XX 1,3-GalT genes. Potential sites to interrupt the alpha-1,3-GalT gene
XX (via homologous recombination) were identified in exons 4, 7, 8 and 9.
XX Such inactivation allows the breeding of 'knock-out' animals. e.g.
XX pigs suitable as donors of organs to overcome hyperacute rejection
XX problems in human xenotransplantation.
XX
XX Sequence 371 AA:
QY 1 MNVCKVILSMVSTVIVFWEYIHSFEGSLFWINPSRNPVCGSSJOKGMILPRWFNN 60
   |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
Db 1 mnvkvivlsmvstvmvfweyihsfegslfwinygsknpv-qssaqrdwipswinn 59
QY 61 G---VHERDGDINFEKEQNRND-ESKILSDMFPNPPKREVVMTMKAPVVECTYNRA 116
   1:||||| 1:||||| 1:|||||:|||||:|||||:|||||:|||||:|||||
Db 60 gthsyhecdadagnekqrkednrelplvdfotpepckrpevntltmkapvwegtyara 119
QY 117 VLDNYAMOKITVCLTAVAGRTIEHYILEEFJLSANKHFWGCHPVIPIYIMVDVSRMPLI 176

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DB 120 vldnyakrkltvgtvldavqylychyleelllsanlylmvqhkvtlyimvddlsmp11 179
 QY 177 ELGPIRSFKVFKIKPEKQWQDLSNMHMKTTGCHIVAHIOHPDPLFCMDVQVQKFCV 236
 DB 180 elqprstskvteokskcwqdiismminkllyobllldhqlhevdlldmvdqylyimvddlsmp11 239
 QY 247 ETLDFSVAGLQAWMYKALDNPDTTEREKESAAVTPFGSGDYVYAAIIGCTPTGVANTU 296
 DB 240 elqpsvqqlqawmykaldpdttyctkesaaylptqpdlyyhaa119qplqylnltq 299
 QY 297 ECFKGLTKOKKNDLEAQMDESHLNKPYFLNKPKTLSPFYCWYHIGLPAIDIKLVKNSW 356
 DB 400 ectrkqllqkndleawmdeshlnkpylltkprklispycwylhigmsvdlitvklaw 359
 QY 457 QIKRYNVVNNV 368
 DB 460 qkkyulvtrml 371

RESULT 5

AA062508
 ID AA062508 standard; protein: 459 AA.

AC AA062508;

PT 26 JUN-1995 (first entry)

DE Galactosyl transferase clone product.

XX gal-alpha (1,3) galactosyl transferase; xenograft; transplant;

KW refection.

OS Sus scrofa domestica.

XX W09421799 A.

PN W09421799 A.

PD 2/9 SEP-1994.

PE 15 MAR-1994; 94AWO A000126.

PK 15 MAR-1994; 94AD 0007H54.

PA (AUST) AUSTIN RES INST.

PI McKenzie IFC, Sandrin MS;

XX WPI: 1994 417019/39.

XX N-PSDB: AAG74712.

PA DNA sequences encoding gal-alpha (1,3)galactosyl transferase

PT and clones family. Such sequences are used in xenograft therapies

XX Disclousure: Page 59; 56pp; English.

PS The sequence is that of the product of the porcine Gal-alpha (1,3)

CC galactosyl transferase gene which produces a Gal epitope on the

CC surface of porcine cells. This epitope is recognised by antibodies

CC which are responsible for hyperacute rejection of xenotransplanted

CC pig cells, tissues and organs.

CC See also AA062507.

CC Sequence: 459 AA;

QY Query Match 62.3%; Score 1648.5; ID 15; Length 359;

DB Best Local Similarity 61.5%; Ident. No. 176-158;

DB Mismatches 303; Conservative 30; Mismatches 22; Indels 17; Gaps 4;

QY 61 G---VHEEKGIDINEEKDORND--ESKLIKLSLQNFNKKRPVNTTKKALVNVKCTYNKA 116
 DB 48 qlshyheevdeatigqekqkrkdnrtqqlplvdmtpokrpvvlitwakpavwqlytra 107
 QY 117 VLIDNYAKOKITVGLIYAVAVRYTEHLEPFTLSANKHFWGHHVLYFYIWDVSRMPL 176
 DB 108 vldnyakrkltvgtvldavqylychyleelllsanlylmvqhkvtlyimvddlsmp11 167
 QY 177 ELGPIRSFKVFKIKPEKQWQDLSNMHMKTTGCHIVAHIOHPDPLFCMDVQVQKFCV 236
 DB 168 elqprstskvteokskcwqdiismminkllyobllldhqlhevdlldmvdqylyimvddlsmp11 227
 QY 227 ETLDFSVAGLQAWMYKALDNPDTTEREKESAAVTPFGSGDYVYAAIIGCTPTGVANTU 296
 DB 228 elqpsvqqlqawmykaldpdttyctkesaaylptqpdlyyhaa119qplqylnltq 287
 QY 297 ECFKGLTKOKKNDLEAQMDESHLNKPYFLNKPKTLSPFYCWYHIGLPAIDIKLVKNSW 356
 DB 288 ectrkqllqkndleawmdeshlnkpylltkprklispycwylhigmsvdlitvklaw 347
 QY 457 QIKRYNVVNNV 368
 DB 448 qkkyulvtrml 359

RESULT 6

AA090573
 ID AA090573 standard; protein: 359 AA.

AC AA090573;

PT 08 APR-1996 (first entry)

DE Pig alpha(1,3)-galactosyltransferase.

XX Alpha(1,3)galactosyltransferase; xenograft hyperacute rejection;

KW Transplantation; galactose alpha(1,3) galactose.

OS Sus scrofa.

XX W09534202-A1.

PN W09534202-A1.

PD 21-DEC-1995.

PE 14-JUN-1995; 95MO-0S07554.

PK 21-JUN-1994; 94US-0278282.

PA 15-JUN-1994; 94US-0260201.

PA (ALEX) ALEXION PHARM INC.

PA (AUST-) AUSTIN RES INST.

PI Fodor WJ, McKenzie IFC, Kotler RP, Sandrin MS, Squinto SP;

XX WPI: 1996-04926/05.

XX N-PSDB: AAT12242.

XX Redn. of rejection of xenogeneic cells following transplantation

PT by introducing a vector expressing fucosyl:transferase into the

CC cells

CC Example 2: Page 52-54; 69pp; English.

CC Pig alpha(1,3)-galactosyltransferase (AA090573) was expressed in

CC monkey COS cells following transfection of the cells with

CC vector pGT which contains an insert including the encoding

CC cDNA (AAT12242). Co-transfection of these cells with vector pHT

CC encoding human H-transferase (AA090572) resulted in a reduction in

CC the levels of galactose alpha(1,3) galactose epitopes expressed

CC by the cells.

CC Sequence: 359 AA;

Query Match 82.1%: Score 1648.5; DB 17; Length 359;
 Best local Similarity 81.5%: Pred. No. 1.7e-158;
 Matches 303; Conservative 30; Mismatches 22; Indels 17; Gaps 4;

QY 1 MNVKKVILSMVSTVIVFWEYHSPEGSLEFWINPSRNPVCGSSLOKGMWLPFRFNN 60
 DB 1 mrvkqvvlsmllvstvmvltweyl-----nrpev-qssaqrgwfwpswin 47
 QY 61 G---YHEEDGDI NEKEQBNED-ESKILSLDFNPPKPEVVTMTKKAIVVWEGCTYNRA 116
 DB 48 qhsyhceedaignekqekednrgelplvdwlnpekrpevvltrwkapvwegtyra 107
 QY 117 VLDNYAAKQITVGLTAVAGRYTEHYLEEFELTSANKRHPVGFYIMVDVSRMPLI 176
 DB 108 vldnyyakqitvgtltavagrytehylectlsanrlymvgkhvlfylmvdvdsrmpil 167
 QY 177 ELGPLRSFKVFKIKPEKKMODISMKMKITIGEHIVAHIOHEVDPLFCMDVQVDFKFGV 236
 DB 168 elgplrstfkvfkikpekkgwgdismmrmtkigehilahqhevdlifclvdvqvtgnitgv 227
 QY 237 ETLGESAQIQAMWYKADPNDFTYERKESAAVTPREGDFFVYAAIFGCTPTQVLTNTQ 296
 DB 228 etlgsqvagiqawwykahpdeftyerkesaaylplfgqgdtyyhaalqtrptqvlntq 287
 QY 297 ECFKGLDKKKNDIEAQWHDESHLLNKYFLNKPRTKILSPYCWMDHYHGLPADIKLVKMSW 356
 DB 288 ecfkglldkknndieawhdeshllnkylflnkpklispycwmdyhmvsdvdrivklaw 347
 QY 357 QTKEXVNVVNNV 368
 DB 348 qtkexvnlvrnvl 359

RESULT 7
 AAM49687
 ID AAM49687 standard: Protein: 363 AA.
 XX AAM49687:
 AC AAM49687:
 XX
 DI 10-NOV-1998 (first entry)
 XX
 DE Porcine alpha-1,3-galactosyl transferase isoform 2.
 XX
 KM Isoform: porcine; enzyme: alpha-1,3-galactosyl transferase; galactose;
 KM sugar: N-acetylglucosamine; glycoprotein: glycolipid; antibody: pig;
 KM graft tissue rejection; organ transplantation; xenotransplant.
 XX
 OS Sus scrofa.
 PN FR2751346-A1.
 XX
 PD 23-JAN-1998.
 XX
 PE 19-JUL-1996; 96FR-0009077.
 XX
 PR 19-JUL-1996; 96FR-0009077.
 XX
 PA (INRM) INSERM INST NAT SANTE & RECH MEDICALE.
 XX
 PI Pourcel C, Souillion JP, Vanhove B;
 XX
 OS WPI: 1998-112876/11.
 DR N-PSDB: AAV49454.
 XX
 PT Transgenic non-human donors of organs for human recipients -
 PT containing DNA encoding antibodies that inhibit graft rejection
 XX
 PS Claim 4: Page 36-37; 71pp; French.
 CC This sequence represents isoform 2 of the porcine enzyme
 CC alpha-1,3-galactosyl transferase (alpha-1,3-GT). The enzyme catalyses
 CC the attachment of a galactose sugar molecule on the N-acetylglucosamine

CC moiety found on surface glycoproteins and glycolipids. These sugar
 CC molecules are partly responsible for raising anti-graft antibodies, which
 CC lead to graft tissue rejection. The invention relates to a method of
 CC inhibiting the graft rejection mechanism by introducing the sequence
 CC encoding an antibody targeted to alpha-1,3-GT into the cells of animal,
 CC especially a pig, from whom organs may be used for xenotransplants.
 CC Neutralisation of the alpha-1,3-GT leads to tissues or organs lacking
 CC the galactose on the glycoproteins and glycolipids, thus preventing
 CC induction of the rejection response.

SO Sequence 363 AA;

Query Match 82.0%: Score 1646.5; DB 19; Length 363;
 Best local Similarity 81.5%: Pred. No. 2.8e-158;
 Matches 303; Conservative 29; Mismatches 23; Indels 17; Gaps 4;

QY 1 MNVKKVILSMVSTVIVFWEYHSPEGSLEFWINPSRNPVCGSSLOKGMWLPFRFNN 60
 DB 5 mrvkqvvlsmllvstvmvltweyl-----nrpev-qssaqrgwfwpswin 51
 QY 61 G---YHEEDGDI NEKEQBNED-ESKILSLDFNPPKPEVVTMTKKAIVVWEGCTYNRA 116
 DB 52 qhsyhceedaignekqekednrgelplvdwlnpekrpevvltrwkapvwegtyra 111
 QY 117 VLDNYAAKQITVGLTAVAGRYTEHYLEEFELTSANKRHPVGFYIMVDVSRMPLI 176
 DB 112 vldnyyakqitvgtltavagrytehylectlsanrlymvgkhvlfylmvdvdsrmpil 171
 QY 177 ELGPLRSFKVFKIKPEKKMODISMKMKITIGEHIVAHIOHEVDPLFCMDVQVDFKFGV 236
 DB 172 elgplrstfkvfkikpekkgwgdismmrmtkigehilahqhevdlifclvdvqvtgnitgv 231
 QY 237 ETLGESAQIQAMWYKADPNDFTYERKESAAVTPREGDFFVYAAIFGCTPTQVLTNTQ 296
 DB 232 etlgsqvagiqawwykahpdeftyerkesaaylplfgqgdtyyhaalqtrptqvlntq 291
 QY 297 ECFKGLDKKKNDIEAQWHDESHLLNKYFLNKPRTKILSPYCWMDHYHGLPADIKLVKMSW 356
 DB 292 ecfkglldkknndieawhdeshllnkylflnkpklispycwmdyhmvsdvdrivklaw 351
 QY 357 QTKEXVNVVNNV 368
 DB 352 qtkexvnlvrnvl 363

RESULT 8
 AAM49688
 ID AAM49688 standard: Protein: 354 AA.
 XX AAM49688:
 AC AAM49688:
 XX
 DI 10-NOV-1998 (first entry)
 XX
 DE Porcine alpha-1,3-galactosyl transferase isoform 3.
 XX
 KM Isoform: porcine; enzyme: alpha-1,3-galactosyl transferase; galactose;
 KM sugar: N-acetylglucosamine; glycoprotein: glycolipid; antibody: pig;
 KM graft tissue rejection; organ transplantation; xenotransplant.
 XX
 OS Sus scrofa.
 PN FR2751346-A1.
 XX
 PD 23-JAN-1998.
 XX
 PE 19-JUL-1996; 96FR-0009077.
 XX
 PR 19-JUL-1996; 96FR-0009077.
 XX
 PA (INRM) INSERM INST NAT SANTE & RECH MEDICALE.
 XX
 PI Pourcel C, Souillion JP, Vanhove B;

```

XX  WFL: 1998 112876/11.
XX  N-PSDB: AAV49456.
XX  Transgenic non human donors of organs for human recipients -
XX  containing DNA encoding antibodies that inhibit graft rejection
XX
XX  Claim 4: Page 43-44: 71pp; French.
XX
XX  This sequence represents isoform 3 of the porcine enzyme
XX  alpha-1,3-galactosyl transferase (alpha-1,3-GT). The enzyme catalyses
XX  the attachment of a galactose sugar molecule on the N-acetylglucosamine
XX  moiety found on surface glycoproteins and glycolipids. These sugar
XX  molecules are partly responsible for raising anti-graft antibodies, which
XX  lead to graft tissue rejection. The invention relates to a method of
XX  inhibiting the graft rejection mechanism by introducing the sequence
XX  encoding an antibody targeted to alpha-1,3-GT into the cells of animal,
XX  especially a pig, from whom organs may be used for xenotransplants.
XX  Neutralisation of the alpha-1,3-GT leads to tissues of organs lacking
XX  the galactose on the glycoproteins and glycolipids, thus preventing
XX  induction of the rejection response.
XX
XX  Sequence: 454 AA.
XX
XX  Query Match: 79.98; Score 1604; DB 19; Length 354;
XX  Best Local Similarity: 79.98; Pred. No. 5,40-154;
XX  Matches: 295; Conservative: 40; Mismatches: 24; Indels: 20; Gaps: 2;
XX
XX  1 MNVKGKYLMLVSVIVVFWYHSHSPGSLFWINPSNRPVGGSSLOKQWMLRWKFN 60
XX  |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:
XX  5 mrvkgvvlsmllvstvmvvtwvynpsgslfwyqsk-----th 45
XX
XX  61 GYDEEDDINFEKEQNEDE-ESKRLKSLDNEENPKREKVYMTKWKAPVWVESTYKAV 119
XX  |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX  46 sylveceatqnekeqkcdarqcelplvdrwmpokrpvevltlwkapvwvqdyrnavid 105
XX
XX  20 NYAKOKLTVGLTVFVAVGKYLEYLEETTSANKRPVGNRPVLYIMVWVSKMPLLEIG 179
XX  |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX  106 nyakokltlvgltdvgyrlyehyleetltsantymvghkvlylmvddlstmpllleig 165
XX
XX  180 FLSKFKVYKTRPKRWQDISMNRKKTIGETVAHGDHEVETLFCMIVGVYQDKKSVET 239
XX  |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX  106 flskfkvylsmllvstvmvvtwvynpsgslfwyqsk-----th 45
XX
XX  240 GESVAULQAMWYKADPNDFTEYERKESAAVLPFGEGDYTHAAIFGCTPTQVINTQRF 299
XX  |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX  106 gsvaulqamwykadhpdftetyerkesaavlpfggdythaaifgctptqyintqrf 285
XX
XX  400 KGLTKRKNDEQWMDHSHLNKYFLINKPTKLLSPGYCWYHIGLPADIKLVKMSWQTK 359
XX  |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX  286 kgltkrkndeqwmdshlnkyflinkptkllspgycwdyhiglpadiklvkmswtk 345
XX
XX  460 EYNNVNNV 368
XX  |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX  446 eynnvnnv 354
XX
XX  RESULT 9
XX  ID AAM49689 standard: Protein: 342 AA.
XX  AC AAM49689;
XX  DT 10 NOV 1998 (first entry)
XX  DE Porcine alpha-1,3 galactosyl transferase isoform 4.
XX  KM Isoform: porcine; enzyme: alpha-1,3-galactosyl transferase; galactose;
XX  KM sugar; N-acetylglucosamine; glycoprotein; glycolipid; antibody; pig;
XX  KM graft tissue rejection; organ transplantation; xenotransplant.
XX  OS Sus scrofa.

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XX  FR275146-A1.
XX  23-JAN-1998.
XX  19-JUL-1996; 96FR-0009077.
XX  19-JUL-1996; 96FR-0009077.
XX
XX  (INNM) INSERM INST NAT SANTE & RECH MEDICALE.
XX  Poullet C, Souillou JP, Vanhove B;
XX  WFL: 1998 112876/11.
XX  N-PSDB: AAV49456.
XX
XX  Transgenic non human donors of organs for human recipients -
XX  containing DNA encoding antibodies that inhibit graft rejection
XX
XX  Claim 4: Page 43-44: 71pp; French.
XX
XX  This sequence represents isoform 4 of the porcine enzyme
XX  alpha-1,3-galactosyl transferase (alpha-1,3-GT). The enzyme catalyses
XX  the attachment of a galactose sugar molecule on the N-acetylglucosamine
XX  moiety found on surface glycoproteins and glycolipids. These sugar
XX  molecules are partly responsible for raising anti-graft antibodies, which
XX  lead to graft tissue rejection. The invention relates to a method of
XX  inhibiting the graft rejection mechanism by introducing the sequence
XX  encoding an antibody targeted to alpha-1,3-GT into the cells of animal,
XX  especially a pig, from whom organs may be used for xenotransplants.
XX  Neutralisation of the alpha-1,3-GT leads to tissues of organs lacking
XX  the galactose on the glycoproteins and glycolipids, thus preventing
XX  induction of the rejection response.
XX
XX  Sequence: 342 AA.
XX
XX  Query Match: 76.88; Score 1543; DB 19; Length 342;
XX  Best Local Similarity: 77.28; Pred. No. 7,70-148;
XX  Matches: 285; Conservative: 29; Mismatches: 23; Indels: 32; Gaps: 2;
XX
XX  1 MNVKGKYLMLVSVIVVFWYHSHSPGSLFWINPSNRPVGGSSLOKQWMLRWKFN 60
XX  |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX  5 mrvkgvvlsmllvstvmvvtwvynpsgslfwyqsk-----th 45
XX
XX  61 GYDEEDDINFEKEQNEDE-ESKRLKSLDNEENPKREKVYMTKWKAPVWVESTYKAV 119
XX  |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX  106 sylveceatqnekeqkcdarqcelplvdrwmpokrpvevltlwkapvwvqdyrnavid 94
XX
XX  120 NYAKOKLTVGLTVFVAVGKYLEYLEETTSANKRPVGNRPVLYIMVWVSKMPLLEIG 179
XX  |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX  94 nyakokltlvgltdvgyrlyehyleetltsantymvghkvlylmvddlstmpllleig 153
XX
XX  180 FLSKFKVYKTRPKRWQDISMNRKKTIGETVAHGDHEVETLFCMIVGVYQDKKSVET 239
XX  |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX  154 flskfkvylsmllvstvmvvtwvynpsgslfwyqsk-----th 45
XX
XX  240 GESVAULQAMWYKADPNDFTEYERKESAAVLPFGEGDYTHAAIFGCTPTQVINTQRF 299
XX  |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX  214 gsvaulqamwykadhpdftetyerkesaavlpfggdythaaifgctptqyintqrf 273
XX
XX  400 KGLTKRKNDEQWMDHSHLNKYFLINKPTKLLSPGYCWYHIGLPADIKLVKMSWQTK 359
XX  |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX  274 kgltkrkndeqwmdshlnkyflinkptkllspgycwdyhiglpadiklvkmswtk 333
XX
XX  460 EYNNVNNV 368
XX  |||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||
XX  334 eynnvnnv 342
XX
XX  RESULT 10
XX  ID AAK13750 standard: Protein: 394 AA.

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XX AC AAR13750;
XX DT 07-NOV-1991 (first entry)
XX DE GDP-Fuc:[beta-D-Gal(1,4/1,3)]-D-GlcNAc/(Glc)alpha(1,3/1,4)
XX DE -fucosyltransferase.
XX KW Glycosyltransferase.
XX OS Mus musculus.
XX PN M09112340-A.
XX PD 22-AUG-1991.
XX PF 14-FEB-1991; 91WO-0500899.
XX PR 12-DEC-1990; 90US-0627621.
XX PR 14-FEB-1990; 90US-0479858.
XX PR 14-FEB-1990; 90US-0480133.
XX PA (UNMI ) UNIV OF MICHIGAN.
XX PI Lowe JB.
XX PS WPI: 1991-267151/36.
XX PS N-PSDB: AA013331.
XX PT Isolation of gene conveying post-translational characteristic -
XX PT e.g. the presence of soluble or membrane bound oligo or
XX PT polysaccharide or glycosyltransferase.
XX PS Disclosure: Fig 2; 155pp; English.
XX CC The amino acid sequence codes for a protein capable of functioning
XX CC as a UDP-Gal:[beta-D-Gal(1,4)]-D-GlcNAc alpha (1,3)galacto-
XX CC syltransferase. The products of this enzyme, sub-terminal alpha
XX CC (1,3) and alpha(1,4) fucose residues are used in the post-
XX CC translational modification of the oligosaccharides on cell-surface,
XX CC intracellular or secreted proteins or lipids. These can be used for
XX CC the prodn. of diagnostics and therapeutics. There is a single
XX CC transmembrane domain consisting of a 19 amino acid hydrophobic
XX CC segment flanked by basic residues and a large (presumably
XX CC catalytic) C-terminal domain that would ultimately be targeted to
XX CC the lumen of the Golgi. It has two potential N-glycosylation sites
XX CC indicating that as with other glycosyltransferases, it may be
XX CC synthesised as a glycoprotein. It is representative of a Type II
XX CC transmembrane protein. See also AAR13749-R13752.
XX SO Sequence 394 AA:

Query Match 73.5%; Score 1476.5; DB 12; Length 394;
Best Local Similarity 72.2%; Pred. No. 5,2e-141;
Matches 268; Conservative 41; Mismatches 47; Indels 15; Gaps 3.

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QY 238 TLGSEVAQIQAMWYKADPNDFYERREKESAAVTPPEGDEYFHAHFGTPTVLNITOE 297
DB 264 LIQGLVAGIQAWYKASPEKTYEREISAAYIPFGQADLYYHAAI199APDHLNITRE 324
QY 298 CPKGLDKRKNMDIEAOWDESHLNKRYFLINKPTKILSPFYCWYHIGLPADIKLKSQW 357
DB 324 CFKGLIQDKKHDAQWDESHLNKRYFLINKPTKILSPFYCWYHIGLPADIKLKSQW 384
QY 358 TKEYNVVRNNV 368
DB 384 LKEYNLVTRNNV 394

RESULT 11
AAR45935
ID AAR45935 standard; Protein: 394 AA.
XX AC AAR45935;
XX DT 26-JUL-1994 (first entry)
XX DE A glycosyltransferase.
XX DE Glycosyltransferase; fucosyltransferase; GDP-Fuc; in vitro; cell;
XX DE surface; oligosaccharide.
XX OS Homo sapiens.
XX PN W09402616-A.
XX PD 03-FEB-1994.
XX PF 20-JUL-1993; 93WO-US06703.
XX PR 20-JUL-1992; 92US-0914281.
XX PA (UNMI ) UNIV MICHIGAN.
XX PI Lowe JB.
XX PS WPI: 1994-048874/06.
XX PS N-PSDB: AA056907.
XX PT DNA fragment encoding a glycosyltransferase - can be used for in
XX PT vitro reactions to modify cell surface oligosaccharide(s) e.g.
XX PT blood gp. determinants, to protect against transplant rejection
XX PS Disclosure: Fig 2; 249pp; English.
XX CC The sequence is that of a human glycosyl transferase. The enzyme
XX CC may be non glycosylated. This prevents premature loss of enzyme
XX CC activity. It can also be used in in vitro reactions to modify cell
XX CC surface oligosaccharide moets. e.g. blood group determinants.
XX CC See also AAR45933-9.
XX SO Sequence 394 AA:

Query Match 73.5%; Score 1476.5; DB 15; Length 394;
Best Local Similarity 72.2%; Pred. No. 5,2e-141;
Matches 268; Conservative 41; Mismatches 47; Indels 15; Gaps 3.

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Query Match	66.08%	Score 1326	DB 15	Length 313
Best Local Similarity	84.7%	Prod. No. 6.4e-126		
Matches 244	Conservative 22	Mismatches 22	Indels 0	Gaps
QY	81	SKLRLSDLENFPPKPEVYVMTKMKAPVWEGSTYRRAVLIDNYAKOKITVGTTPAVGRYI	14	
Db	26	skalltlticltivrltkpvevcltrwkapvweglytravldnyaqkltvgtltvavgryl	85	
QY	141	EHYLEEFLLTSANKKIPMGCHPVLYIYVDDVSRLPLIELGLRSKRVKIKPEKRMODISM	200	
Db	86	ehyleeelltsantylmvgkvvilylmvddlsrmllelglrtskvfeikeekrwqdlsm	144	
QY	201	MRMKTIGHIVAHVQHVKDPLFCMDVQVYFQDKRGVETLGSVAGQLOAMWYKADPNDFTY	266	
Db	206	erpkesaaylprfgdlyhyaalltggdrlqylntlqecfkrlldqkendlaaewndesyl	266	
QY	321	NKRYLNNKPKILSPFYGMVDYHIGLPADIKLVKMSWQTLREYNNVNNY	368	
Db	266	nkyllnnkpkilspcywdyhlqmsvdrlrvqagwqkkeynlvrnl	313	
RESULT 14				
AAK57021	ID	AAK57021 standard; Protein: 375 AA.		
AAK57021	AC			
AAK57021	XX			
DT	16	FEB-1995 (first entry)		
XX	DE	Human A transferase (A2 subtype).		
XX	DE	Blood; group; determinant; antigen; erythrocyte; oligosaccharide;		
KW	glycoconjugate; glycosphingolipid; glycoprotein; glycosyltransferase.			
KW	transferase.			
XX	XX			
OS	Homo sapiens.			
XX	XX			
XX	XX			
FT	Key	Location/Qualifiers		
FT	Region	1..53		
FT	/label=	Unsure.		
FT	/note=	"These amino acids are unknown."		
PN	055326857-A.			
XX	XX			
FD	05-JUL-1994.			
XX	XX			
XX	PF	31-AUG-1989; 8905-0402695.		
XX	PR	31-AUG-1989; 8905-0402695.		
XX	PR	29-AUG-1991; 9105-0752101.		
XX	FA	(BIOM-.) BIOMEMBRANE INST.		
PI	Clausen H, Hakomori S, White T, Yamamoto F;			
XX	XX			
DR	WPI: 1994-217096/26.			
XX	XX			
PT	Isolated DNA molecules - encode human histo-blood groups A, B,			
XX	and O-glycotransferases			
XX	XX			
PS	Example 8: Figure 7: 63pp; English.			
XX	XX			
CC	The histo-blood group ABH determinants are major allogeneic antigens			
CC	in both erythrocytes and tissues of humans. They generally			
CC	constitute peripheral parts of the oligosaccharide chains of			
CC	glycoconjugates i.e. linked to lipids (glycosphingolipids) or to			
CC	proteins (glycoproteins). It was proposed that the A and B			
CC	phenotypes were associated with glycosyltransferases that converted			
CC	the H substance associated with the O phenotype to A and B			

[illegible]

A hydrophobic region spanning 21 amino acids precedes this N terminus and appears to be the transmembrane region of the membrane-bound form of A' transase. A proline-rich region (by out of 60) follows the hydrophobic region. An N-glycosylation site appears to be located at position 112 (N-F-T). The remaining long C-terminal portion is moderately hydrophilic. See also A0011489 and A0R1786-93.

XX	353 AA;
50	Sequencia

Query Match 32.78; Score 656; DB 12; Length 353;

Best Local Similarity 49.18; Prod. NO. 5.56-58.2
Matches 125; Conservative 51; Mismatches 97

Matches 125; conservative 51; mismatches 97; indels 4; gaps 32

[illegible]

Search completed. November 5, 2001, 15:20:31
Job time: 8519 sec

Job time: 8519 sec.